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GNE 3030R1C1

PATENT

**IN THE UNITED STATES PATENT AND TRADEMARK OFFICE**

Applicant	:	Desnoyers, et al.
Appl. No.	:	09/931,836
Filed	:	August 16, 2001
For	:	NOVEL PEPTIDES THAT INDUCE CHONDROCYTE REDIFFERENTIATION
Examiner	:	Jiang, Dong
Group Art Unit	:	1646

**DECLARATION OF LUC DESNOYERS AND WILLIAM I. WOOD**  
**UNDER 37 CFR §1.131**

Commissioner for Patents  
P.O. Box 1450  
Alexandria, VA 22313-1450

Dear Sir:

We, Luc Desnoyers and William I. Wood, declare and state as follows:

1. We are the inventors of the subject matter that is presently claimed in the above-captioned patent application.
2. During the time period in which all of the events and activities described herein occurred, we were employed by Genentech, Inc., the assignee of the above-captioned application.
3. All of the events and activities described herein were performed by us personally, or under our direction, as part of our duties as employees of Genentech, Inc.
4. The invention claimed in the above-captioned patent application was conceived prior to April 20, 1999 and diligently reduced to practice thereafter in the U.S. as described below.
5. Prior to April 20, 1999, we conceived of the polypeptides claimed in the above-captioned patent application. This is demonstrated by the attached sequence printout (Exhibit A), which was generated prior to April 20, 1999, and which shows the complete sequence of the polypeptide having the sequence of SEQ ID NO:2. The attached printout also shows the complete sequence of the nucleic acid which has the sequence of SEQ ID NO:1. As evidenced by the sequence printout, we were in possession of the complete polypeptide sequence prior to April 20, 1999.
6. The date deleted from page 1 of Exhibit A is a date prior to April 20, 1999, and was redacted pursuant to M.P.E.P. § 715.07. The redacted date is the date when the data were generated; the date the report was printed, April 16, 2004, remains on the report.

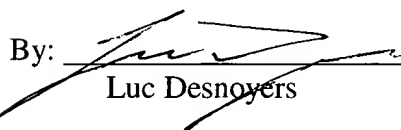
Appl. No. : 09/931,836  
Filed : August 16, 2001

7. After initially conceiving the polypeptide having the sequence of SEQ ID NO:2 prior to April 20, 1999, we diligently reduced the claimed subject matter to practice by working to express and purify the polypeptide and to run it systematically through many assays. The cDNA was deposited with the American Type Culture Collection (ATCC) on January 12, 1999 and assigned ATCC no. 203581. The protein of interest was assigned a "protein inventory number" (e.g., PIN1308 and PIN1308-1). As set forth in the enclosed Exhibit B, the polypeptide was expressed in *E. coli* - PUR1009 (see page 2) on November 16, 1998; in *Baculovirus* - PUR1039 (see page 3) on November 23, 1998; and in mammalian cells (see page 4) on February 17, 1999. Furthermore, various constructs with poly-His or IgG tags were made from the time of first cloning and the construction of these was followed by expression and purification of the protein during the time period of prior to April 20, 1999 through March 13, 2003. For example, Exhibit C shows July 13, 1999 as the date of purification of a polypeptide having the sequence of SEQ ID NO:2. PIN1308 and/or PIN1308-1 were distributed to various scientists for multiple cell-based assays and/or quality confirmation tests from August 20, 1999 through January 22, 2001.

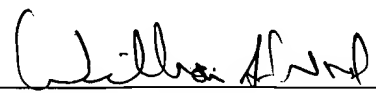
8. Exhibits D and E list the assays performed on the purified protein. Assay ASY110, called "Chondrocyte Re-differentiation Assay" was completed on November 10, 1999 for PIN1308-1, which is a polypeptide having the sequence of SEQ ID NO:2. PIN1308-1 was delivered to Luc Desnoyers for one of the assay runs on October 22, 1999; testing was completed on November 10, 1999. Exhibit E is an assay result list that shows positive results for the assay completed on November 10, 1999, thereby confirming the ability of the claimed polypeptide to induce chondrocyte redifferentiation. Thus, actual reduction to practice occurred at least by November 10, 1999.

9. After reducing the invention to practice, we worked with the Genentech, Inc. patent department to prepare a non-provisional patent application, which included the sequence of SEQ ID NO:2, as well as the data showing the ability to induce chondrocyte redifferentiation. That application was filed on March 1, 2000.

10. We hereby declare that all statements made herein of our own knowledge are true and that all statements made on information or belief are believed to be true, and further that these statements were made with the knowledge that willful false statements and the like so made are punishable by fine or imprisonment, or both, under Section 1001 of Title 18 of the United States Code and that such willful statements may jeopardize the validity of the application or any patent issued thereon.

By:   
Luc Desnoyers

Date: 5/17/04

By:   
William I. Wood

Date: 8/17/04

**Appl. No.** : **09/931,836**  
**Filed** : **August 16, 2001**

## **EXHIBIT A**

(16 pages; pages 4-19)

EXHIBIT A—PAGE 1

>Friday, April 16, 2004

>DNA44686 [Full]

>584 Sites [All Sites]

> [DNA44686, sheldens

>Sequence confirmed by parap.

```

      rnaI
sau3AI naeI
mboI/ndelI[dam-]
      dnlI[dam-]
      dnlI[dam+]
      alwI[dam-] sau3AI
      nlaIV xbaI mboI/ndeI[dam-]
      haeIII/palI bfaI dnlI[dam-]
      mmoI haeIII/palI bfaI dnlI[dam-]
      bglI[M.haeIII-] hpy88III taqI
      tfII apoI sfiI eaeI bstYI/xhoII dnlI[dam+]
      hinfI[M.taqI-] cfrI bamHI[M.mspI-] mnlI
      taqI[M.claI-] haeIII/palI alwI[dam-] alwI[dam-]
      claI/bsp106 eaeI bsrI mspI[M.bamHI-][M.haeIII-] taqI
      bspDI[dam-] cfrI tpeRI hpaII mnlI bstYI/xhoII mnlI dndI bsrKI
      1 CAACTGCACC TCGGTCTAT CGATTCGAT TCGGCACAC TGGCCGAT CCTCGACCT CGACCCAGC GTCCGGCAT CTGCCGAGG
      TTGACCTGG AGCCACATA GCTAAGCTTA AGCCGCTGTG ACCGCGCTAG CAGATCTCTA GGGAGCTGGA GCTGGGTGCG CAGGCCGTA GACGGGCTCC
      insert starts here

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GSeqEdit, DNA44686 [Full], page 1

EXHIBIT A—PAGE 2

```

sstI
sacI
HgiAI/aspHI (M.aluI-)
ec1136I-
bsp1286 (M.aluI-)
bsiHXAI
bmy-
banII (M.aluI-)
scrFI (dcn-)
pspGI
mvaI aluI
ecoRII (dcn-)
dsav (dcn-)
bstXI
bssKI (dcn-)
mwoI bptII/gauI (dcn-)
bstXI apyI (dcn+)
101 AGACCAACGCT CCTGGAGCTC TCCTGCTCTC TCAGGGAGAC TCTGAGGCTC TGTGAGAGAT CATGCTTGG AGGCACTCA TCTATTGGCA ACIGCTGGCT
TCTGCTGCGA GGACTTCGAG ACACAGAG AGTCCCTCTG AGACTCCGAG ACACACTCTTA GTACGAAACC TCCGTCGACT AGATAACCT TGACGACCGA
1
N L W R Q L I Y W Q L L A
*MET
pleI mnlI
nlyI mspI hpyCHAV
hinfI bslI hpaII sfiI bsmAI
nmlI hpyI8III nlaIII bsaNI pstI nlaIII sfc
20. TGTGTTTCC TCGCTTTTC CCTGTGICAA GATGATACA TGGAGTCTCC ACACACCGGA GGACTACCCC CAGACTGCGAG TAACTGTGT CATGGAGCT
AACAAAAAGG AGGAAAAAC GGACACAGTT CTACTATGC ACCTCAGAGG TGTIIGCCCT CCTGATGGGG GTCTGACGTC ATTACACACA GTACCTCTGA
14 L F F L F C L C Q D E Y N E S P Q T G G L P P D C S K C C H G D Y

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EXHIBIT A—PAGE 3

```
mwol
  bglI[M.haeIII-]
  sau96I[M.haeIII-]
  sau96I[M.haeIII-]
  pspOMI/bsp120I
  nlaIV scrFI[dcM-]
    sau96I[dcM-][M.haeIII-]
    scrFI[dcM-] eco0109I/draII scrFI[dcM-]
    pspGI scrFI[M.hpall-] pspGI
    mvaI nciI pspGI mvaI nlaIII
    ecoRII[dcM-] haeIII/paII ecoRII[dcM-]
    dsav[dcM-] bsp1286[M.haeIII-] xcmI
    bstNI nspI mvaI dsav[dcM-]
    bssKI[dcM-] bmyI ecoRII[dcM-] styI
    bsaJI hpall dsav[dcM-] ncoI
    sau96I[M.haeIII-] banII[M.haeIII-] bstNI dsal
    xcmI nlaIV apy-[dcM+] apaI bstNI bssKI[dcM-]
    styI haeIII/pa-I dsav bssKI[dcM-] bsgI/bstDSI
    mwol mnli bsaJI bsaJ haeIII/paII bsaKI mnli bsmI apyI[dcM+]
    aluI tagI mwol eco0109I/draII bsaKI mnli bsmI apyI[dcM+]
301 ACAGCTTTCG AGCTACCAA GGCOCCTCTG GGCACCGGG CCTCTCTGGC ATTCAGGAA ACCATGGAAC CAATGCAAC AATGAGCCA CTGGTCATCA
TCTCGAAGAC TCCCAIGGVT CCGGGGGGAC CCGGTGGCCC GGGAGGACCG TANGTCCTT TGGTACCTTT GTTACCTGGT GACCACTACT
48 S F R C Y Q G P P G P P G P P G I P G N H G N N G N K G A T G H E
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GSeqEdit, DNA4686 [Full], page 3

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EXHIBIT A—PAGE 6

```

nlaIII
styI
ncoI sau96I
dsal nlaIV
tflI
hinfI
hpyCH4V
ddelI
801 AAAGGGGATG AGGTTCGCT GCGAATGGGC AATGGCGCTC TCCATGGGGA CCACCAAGC TTCTCCACT TTCCAGGATT CTTGCTCTTT GAACCTAAGT
TTTCCCTTAC TCCAAACCGA CGCTTACCGG TTACCGCCGAG AGGTACCCCT GGTGGTTGCG AAGAGGTGGA AAGTCTCTAA GSACGAGAAA CTTTGATTCA
214 K G D E V W I R M G N G A L H G D H Q R F S T F A G F L L F E T K O

mnlI
ddeI
bspCNI
sau3AI
mboI/ndeII[dam-]
dpmI{dam-}
trp9I
maeIII hpy188I nseI mnlI bsrDI
901 AAATATATGA CTGCAATAGC TCCACTTGG GGAAGACTTG TAGCTAGCT GATTGTGATC GATCTGAGGA ACATTAAAGT TGAGGGTTTT ACATTGCTGT
TTATATATCT GATCTATCG AGGTGNAACC CTTCTGNAAC ATGCACTCGA CTAACAATG CTAGACTCCT TGTAAATTCA ACTCCCAAA TGTACGACAA

ddelI
bsp1286 tflI dd
bmyI hpy188I mboII
banII mboII bpuAI
tsp509I bspCNI hinfI bbsI bs
bsrDI hpyCH4V sfiI csp6I
tsp509I hpyCH4V
1001 ACTCAAAAAT TATTTGGTTG CAATGTTGTC CAGCTACAG GTACACCAAT AATGTTGGAC ATTCAGGGG CTCAGAGAA TCACCCACAA AATAGTCTTC
TAAGTTCTTT AATACCAAS GTACACAA GCGGATGTC CAATGTTGTA TTACAACTG TTAAGTCTTC GAGTCTCTTT AGTTGTTCTT TATCAGAGG

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GSeqEdit, DNA4686 [Full], page 6

EXHIBIT A—PAGE 7

nlaIV  
 bsrI  
 styI  
 bsaJI  
 mnlI  
 bseRI  
 hgaI  
 bspMI  
 hpy188I  
 tsp509I  
 hpy188I  
 sspI  
 1101 TCAGTGGAC TTGACTAATA TCTCAGCAT CTTATCACT CTTTCCTGG CACCTAAAG ATAAATTCCTC TCTGACGCG AGTGGAAATA TTTTCTTA  
 AGCTACTCG AACTGATTAT ACGAGICGTA GAAATAGTGA GMAAGGACG GTGATTTC TANTAGAGG AGACTGGTC CAPCCTTAT AAAAAGAGAC  
 tru9I  
 tep503:  
 apoI  
 hpyCH4V  
 tsp509I  
 mnlI  
 bsrI  
 hpy188III  
 ecc57I  
 mseI  
 1201 TCACAGAGCT CATTTGCARA GAATTTGAC TACTCTGCTT TTAATTAAT ACCAGTTTC AGGAACCCCT GAAGTTTAA GTTCAATTATT CTTTATAACA  
 ACTGTCTCA GTAAAGCTT CTTAATCTG ATGAGACGAA AATTAATTA TGGTCAAG TCCTTGGGGA CTTCAAAAT CAAGTAATAA GAAATATTGT  
 tseI  
 fru4HI/dsoFI  
 rmaI  
 maeI  
 mvoI  
 bbvI  
 bstAPI  
 bsp1286  
 aluI  
 bmyI  
 bfaI  
 mvoI  
 aiul  
 tsp509I  
 mnlI  
 ddeI/M.  
 celII/es  
 blpI/bpu  
 aluI  
 hpy188I  
 tfiI  
 foki  
 hinfI  
 bstFI  
 1301 TTTCAGGAA TCGGATCTAG TGATATGACA GGGCTGGGGC AGAAGACGG GCACTAGCTG CCTTAATAGC TAATTTAGTG CCTCCCTGT TCAGCTTAGC  
 AACTCTCTI AGCTACATC ACTATCTGT CCGACCCCG TTCTTCTCC CGTATCGAC GGAATATCG AATTAATJAC GGGAGGCACA AGTCGAATCG

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EXHIBIT A—PAGE 10

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> length: 1906

accI (GTAKAC) :
acII (CCGC) :
afIII (ACRYGT) :
ahaiI (TTTAA) :
aluI (AGCT) :
a1WI (GGATCANN) :
apaI (GGGCC) :
apoI (RATTY) :
apyI (CCNGG) :
aseI (ATTAT) :
asni (ATTAT) :
aspiI (GWCWC) :
avaI (CYGCG) :
avaII (GWC) :
banHi (GCATCC) :
banI (GGYRC) :
banII (GRCYC) :
bbsI (GAAGACNNNN) :
bbv- (CAEC) :
b-fai (CTAG) :
bglI (GCCNNNGGC) :
blpI (GCTNAGC) :
bnyI (GDCCHC) :
bpmI (CTGAG) :
bpu-102- (GCTNAGC) :
bpuAI (GAAGACNNNN) :

1032
452 1815 1819 :870
77
1464
116 :75 303 741 793 918 942 947 1356 1368 1393 1483 1863 1896
46 47 58 1419
338 628
27 1221 1444
111 327 345 354 434 1713
1683
1683
115
94 442 488
848
46
1149
115 338 628 1068
125 726 932 1095
173 458 818 1357 1894
53 795 911 1354 1827
34 340 1869
943 :394
:15 338 628 1368 1349 1378
:12
943 1394
125 726 932 1095

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EXHIBIT A—PAGE 11

```
bsaI (GGTCTC(NNNNN)) : 100 582
bsaJ- (CCNNGG) : 9 95 317 326 327 362 434 488 489 842 :145 1873
bsaW (NCCGGW) : 255
bseRI (CAGGAGNN(NNNNNNN)) : 97 1167
bsgI (GTGCAG) : 4
bsh.1236I (CGCG) : 78 1820
bsiCI (TTCGA) : 24
bsiEI (CERYCG) : 1816
bsiHKAI (GNCWC) : 115
bsII (CCNNNNNNGG) : 249 633 922 1544 1837
bsmAI (GTCTC) : 100 136 245 295 582
bsnAI (GTCTC) : 100 136 245 295 582
bsnFI (GGGACNN(NNNNNNNNN)) : 847
bsn- (GATGCN) : 349 516
bsnFI (GCNCC) : 173 458 818 1357 1815 1818 1869 1894
bsp106 (ATCGAT) : 19
bsp120I (GGGCCC) : 338 628
bsp1286 (GDGCHC) : 115 338 628 1068 1349 1378
bsp1407I (TGTACA) : 736
bspCN- (CTCAGNN(NNNNNNN)) : 130 142 944 964 1071 1100 1123
bspJI (ATCGAT) : 19
bspHI (TCATGA) : 395 610
bspM- (ACCTGC) : 1177 1836
bsrBI (GAGCGG) : 450
bsrDI (GCAATGNN) : 829 992 1020
bsrGI (TGTACA) : 736
bsrI (ACTCGN) : 39 390 615 633 1252 1500
bsrK- (CCNNGG) : 83 111 327 336 345 354 434 488 489 1713
bst4CI (ACNGI) : 556 723 1615 1729
bstAPI (GCANNNTGC) : 1351
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EXHIBIT A—PAGE 12

bstBI (TTCGAA) :	24
bstDSI (CCRYGG) :	362 842 1873
bs-EII (GGTNACC) :	429
bstF51 (GGATG) :	680 769 806 1313 1553
bstNI (CCWGG) :	111 327 345 354 434 1713
bstUI (CGCG) :	78 1820
bstXI (CCAMNNNNNTGG) :	104 1500
bs-YI (RGATCY) :	46 57
btgI (CCRYGG) :	362 842 1873
bteI (GCAGTGN) :	574
cac8I (CCNNGC) :	194 794
cel-I (GCTNAGC) :	943 1394
cfoI (GCSC) :	835
cfrI (YGGCCR) :	32 41 1816 1867
clal (ATCGAT) :	19
csp6I (GTAC) :	701 737 1041 1613
dde- (CTNAG) :	130 142 895 944 964 1071 1100 1123 1395 1695
dpnI (GATC) :	47 58 961 1419
dpnII (GATC) :	47 58 961 1419
dral (TTTAAA) :	1464
dralI (RGNCCT) :	320 338 437 627 628
drcI (GACNNNNNGTC) :	72 1823
dsal (CCRYGG) :	362 842 1873
dsav (CCNGG) :	83 111 327 336 345 354 434 488 489 1713
eaeI (YGGCCR) :	32 41 1816 1867
eagI (CGECCG) :	1816
ec11361I (GAGCTC) :	115
ec-X- (CGECCG) :	1816
eco57I (CTGAAG) :	507 542 569 659 728 789 1269 1667
ecoNI (CCNNNNNAGG) :	1837

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EXHIBIT C—PAGE 13

eco01091 (RGGCCY) :	320 338 437 627 628
ecoRI (GAATTC) :	27 1444
ecoRII (CCWGG) :	111 327 345 354 434 1713
esp- (GCTNAGC) :	943 1394
fru4HI (GCNGC) :	173 458 818 1357 1815 1818 1869 1894
fruJII (CGCG) :	78 1820
foxI (GGATC) :	680 769 806 1313 1553
gstI (CTGCAG) :	112
haeII (RGGCGY) :	834
haeIII (GGCC) :	33 42 321 331 339 439 465 629 1817 1868 1877
hgaI (GACGC) :	79 1174
hgiAI (GNGCWC) :	115
hhaI (GCSC) :	835
hmpI (GCGC) :	835
hincII (GTYRAC) :	1645 1832
hindII (GTYRAC) :	1645 1832
hirdIII (AAGCTT) :	1862
hirfI (GATC) :	22 138 157 243 494 877 1078 1308 1823 1830
hpaII (CCGG) :	44 83 256 336 489
hpaI (GGTGA) :	411 429 655
hpy188I (TCNGA) :	141 509 551 762 963 1072 1101 1171 1311 1441 1551 1666
hpy188III (TCNNGA) :	52 227 395 610 1259 1563 1826
hpyCH4II (ACNET) :	556 723 1615 1729
hpyCH4V (CGCA) :	5 276 515 709 872 1019 1215 1640 1839 1893
haeI (CTAG) :	53 795 911 1354 1827
haeIII (GTNAC) :	430 956
hboI (GATC) :	47 58 961 1419
hboI- (GAAGA) :	126 568 652 727 932 1075 1096 1869
hcrI (CGRYCC) :	1816
hliI (ACGCGT) :	77

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EXHIBIT C—PAGE 14

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mslV (GAGTCNNNNN) :
mslI (CCTC) :
mslI (TTAA) :
mslI (CAYNNNRTG) :
mspI (CCGG) :
msvA (CCWGG) :
msvI (CGCG) :
msvI (GCKNNNNNGC) :
mslI (CCSGG) :
mslI (CCATGG) :
mslI (GATC) :
mslI (GCTAGC) :
mslI (CATG) :
mslI (GGNNCC) :
mslI (GGGGCCG) :
mspH (RCATGY) :
mspI (RCATGY) :
mspI (CTCGAG) :
mslI (GGCC) :
mslI (GAGTCNNNN) :
mslI (TTATAA) :
mspI (CCCGG) :
mspI (CCWGG) :
mspI (GGGCC) :
mslI (CTSCAG) :
mslI (TCAAGA) :
mslI (CTAG) :
mslI (GTAC) :
mslI (GAGCTC) :
138 243 1823 1830
9 50 62 68 97 144 176 209 259 310 342 441 445 678 687 810 966 982
169 1382 1567 1573 1610 1653
974 1241 1246 1277 1434 1465 1584 1684 1781
675
44 83 256 336 489
111 327 345 354 434 1713
78 1820
34 198 164 304 313 340 452 516 525 733 1351 1360 1869
83 336 488 489
362 842 1873
47 58 961 1419
794
161 239 291 363 396 462 521 611 665 675 734 780 843 1642 1874
46 321 338 384 402 437 465 627 628 629 847 1149 1262
18:5
733 1641
733 1641
442
33 42 321 331 339 439 465 629 1817 1868 1877
138 243 1823 1830
1293 1899
488
111 327 345 354 434 1713
338 628
275 1838
395 610
53 795 911 1354 1627
70: 737 :041 1613
115

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EXHIBIT C—PAGE 15

sali (ETCGAC) :	1832
sau3AI (GATC) :	47 58 961 1419
sau36I (GGNCC) :	321 330 338 339 438 465 628 629 848 1877
sceI (TAGGGATAACAGGGTAAT) :	1844
scrFI (CCNGG) :	83 111 327 336 345 354 434 488 489 1713
sfaNI (GCATC) :	87 1127
sfoI (CTRYAG) :	275 299 1035 1838
sfil (GCCNNNNNGGCC) :	33 1868
sfuI (TTCCGA) :	24
smaI (CCCGGG) :	488
snlI (CTYRAG) :	442
sepI (AATATT) :	1187
sstI (GAGCTC) :	115
styI (CCWNGG) :	317 362 842 1145 1873
taqI (TCGA) :	20 25 64 70 308 443 1833
tfil (GATC) :	22 157 494 877 1078 1308
thai (GGCG) :	78 1820
tlil (CTCGAG) :	442
trr9I (TTAA) :	974 1241 1246 1277 1434 1465 1584 1684 1781
tseI (SCNGC) :	173 458 818 1357 1694
tsp45I (GTSAC) :	430
tsp503I (AATT) :	28 1009 1061 1163 1222 1243 1372 1445 1516
topRI (NNCACTGNN) :	38 389 557 575 1616
vspI (ATTAAT) :	1683
xbaI (TCTAGA) :	52 1826
xcmI (CCANNNNNNNTGG) :	317 362
xhoI (CTCGAG) :	442
xhoII (RGATCY) :	46 57
xmaI (CCCGGG) :	488
xmaIII (CGGCGG) :	1816

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**Filed** : **August 16, 2001**

## **EXHIBIT B**

(4 pages; page 21-24)

Appl. No. : 09/931,836  
 Filed : August 16, 2001

EXHIBIT B—PAGE 1

Protein Request										
PRODUCTION HISTORY										
UNQ 753										
Order Protein	Transfeco DNA	EXP System	Normal Name	PRO	Protein Request	EXP	PUR	PUR Status	PUR Warning	Culture Vial
1. Order	DNA84665	E Coli	Human CTRP3 Poly-His	PRO1825		EXP2247	PUR1009	Done		
2. Order	DNA84665	E Coli	Human CTRP3 Poly-His	PRO1825		EXP2247	PUR4414	Done		
3. Order	DNA87982	Baculovirus	Human CTRP3 IgG	PRO1855		EXP2255	PUR1039	Drop		1
4. Order	DNA102368	Mammalian Stable	Human CTRP3 Poly-His	PRO4365		EXP2794				

EXHIBIT B—PAGE 2

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 Filed : August 16, 2001

EXHIBIT B—PAGE 3

GENEN GENES		SITEMAP		Additional F																																																																																																																																																	
<a href="#">GENE VIEWER</a> <a href="#">GENE</a> <a href="#">FAM</a> <a href="#">MAP</a> <a href="#">GENEUB</a>		<a href="#">End</a> <a href="#">Now</a> <a href="#">Updates</a>																																																																																																																																																			
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[ASY](#) | [DNA](#) | [DOM](#) | [EXP](#) | [FAM](#) | [FLS](#) | [LIB](#) | [LOT](#) | [MAP](#) | [OLI](#) | [PRB](#) | [PRQ](#) | [PUR](#) | [RNA](#) | [SRC](#) | [UNQ](#) | [XPT](#) | [YST](#)  
[Assay Viewer](#) | [Sequence Viewer](#) | [Gene Viewer](#) | [GenenGenes](#) | [SAGE](#)

GenenGenes Feedback



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EXHIBIT B—PAGE 4

GENEN GENES		SITE MAP		Additional f	
<a href="#">GENE VIEWER</a> <a href="#">GENES</a> <a href="#">FAM</a> <a href="#">MAP</a> <a href="#">GENEHUB</a>		<input type="radio"/> Find <input type="radio"/> New <input type="radio"/> Update			
<a href="#">SEQUENCE VIEWER</a> <a href="#">SRC</a> <a href="#">RNA</a> <a href="#">LIB</a> <a href="#">FLS</a> <a href="#">OLI</a>		<input type="text" value="SELECT"/> <input type="button" value="Go"/>			
<a href="#">ASSAY VIEWER</a> <a href="#">DOM</a> <a href="#">EXP</a> <a href="#">PUR</a> <a href="#">LOT</a> <a href="#">ASY</a>					
<b>EXP2794</b>		<a href="#">VIEW DNA</a>		<a href="#">View Protein</a>	
<p>Gene Info: UNQ753 PRO 4365 Human CTRP3 Poly-His TFDNA102368 FLDNA44686</p> <p>EXP Lab Name: sst.44686.H8</p> <p>DNA Lab Name: sst.44686.H8</p> <p>Protein Request ID:</p> <p>System: Mammalian Stable</p> <p>Exp Cell Harvest Date:</p> <p>Control:</p> <p>Fermentation Run ID:</p> <p>Cell Line: CHO</p> <p>Expression Media:</p> <p>Growth Factors: PS24</p> <p>Supplements:</p> <p>Warning:</p> <p>Gels: <u>GEL180</u> GEL181</p> <p>Expressed: FALSE</p> <p>Comments: no band on western</p> <p>Status:</p> <p>Date Entered: February 16, 1999</p> <p>Date Cancelled:</p> <p>Scientist: <u>Lhney Lewis-Steiner</u></p> <p>Notebook: 30966 - 55</p> <p>Protein Lots:</p>					
<p>Harvest Date:</p> <p>Cell Pellet:</p> <p>Cell Banking ID:</p> <p>F Cell Status:</p> <p># Days Incubated:</p> <p>Transfection Date: February 17, 1999</p> <p>Transfect Date:</p> <p>Transfer Volume:</p> <p>Pub(s):</p> <p>Date Complete:</p> <p>Cancel Reason:</p> <p>Status: Drop</p> <p>Storage Location: Crowley Lab</p>					
No LOTs for this EXPression					
<p> <a href="#">ASY</a>   <a href="#">DNA</a>   <a href="#">DOM</a>   <a href="#">EXP</a>   <a href="#">FAM</a>   <a href="#">FLS</a>   <a href="#">LIB</a>   <a href="#">LOT</a>   <a href="#">MAP</a>   <a href="#">OLI</a>   <a href="#">PRB</a>   <a href="#">PRO</a>   <a href="#">PUR</a>   <a href="#">RNA</a>   <a href="#">SRC</a>   <a href="#">UNQ</a>   <a href="#">XPT</a>   <a href="#">YST</a>  <a href="#">Assay Viewer</a>   <a href="#">Sequence Viewer</a>   <a href="#">Gene Viewer</a>   <a href="#">GenenGenes</a>   <a href="#">SAGE</a> </p> <p><a href="#">GenenGenes Feedback</a></p>					

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## **EXHIBIT C**

(2 pages; pages 26-27)

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EXHIBIT C—PAGE 1

GENEN GENES		SITEMAP		Additional F	
GENE VIEWER   GENE   FAM   MAP   GENENUT SEQUENCE VIEWER   DNA   SRC   RNA   LB   FLS   OLI ASSAY VIEWER   SRC   DOM   EXP   PUR   LOT   ASY		SELECT <input type="text"/> GO			
PUR1009		View DNA		View Protein	
Gene Info: UNQ753 PRO 1825 Human CTRP3 Poly-His TF DNA84665 FL DNA44686					
Protein Request ID					
DNA Lab Name		pE44686-1		Protein Form Name	
PUR Name				Control	
Exp'd PUR Date				PUR Date: July 13, 1999	
EXP		EXP2247		No Sequence report available	
Mass Spec				Sequence Info	
Warning				GELS	
Endotoxin Level		6.24 EU/ml		GEL461	
LLS Mol. Mass (g/mol)				AA Analysis (mg/ml)	
Ext. Coef. (mg/ml) (cm)				OD 280	
Prot. Assay/ml				Endotoxin Units/mg Protein	
Reduced SDS-MW		Approx. 31, 55 kDa		Prot. A ppm	
Theoretical MW of ORF #1		26723.56			
Gel Score					
Buffer		1 mM HCl / 0.15 M NaCl / 4% mannitol			
Comments					
Status					
Date Entered		November 16, 1998		PUR Done Date	
Yield Concentration		4752 nM		Yield Volume: 4.5ml	
Date Canceled				Cancel Reason	
Scientist		Corpuz, Racquel		Status: Done	
Delivered To				Origin	
Notebook		32647-8-		Storage Location	
Protein Lots					
OT2552		PIN1308-1		1009	

ASY | DNA | DOM | EXP | FAM | FLS | LB | LOT | MAP | OLI | PRS | PRO | PUR | RNA | SRC | UNQ | XPT | YST  
 Assay Viewer | Sequence Viewer | Gene Viewer | GenenGenes | SAGE

GenenGenes Feedback

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EXHIBIT C—PAGE 2

PUR1009

GENE GENES

Find New Update

Additional Resources:

View DNA ss. View Protein pI. Update Record

Gene Info: UNQ753 PRO 1825 Human CTRP3 Poly-His TE DNA84685 EL DNA44686

Protein Request ID

DNA Lab Name: pE44686-1

PUR Name

Expctd. PUR Date

EXP: EXP2247

Mass Spec

Warning

Endotoxin Level: 6.24 EU/ml

LLS Molar Mass (g/mol)

Ext.Coef. (mg/ml)<sup>-1</sup>(cm)<sup>-1</sup>

Prot A ng/ml

Reduced SDS MW: Approx. 31, 55 kDa

Theoretical MW of ORF#1: 26723.56

Gel Score

Buffer: 1 mM HCl / 0.15 M NaCl / 4% mannitol

Comments

Status

Date Entered: November 16, 1998

Yield Concentration: 4752 nM

Date Canceled

Scientist: Corneuz, Remyel

Delivered To

Notebook: 32647-8-

Protein Lots

LOT2552

PIN1308-1

1009

Protein Formalname: Control

PUR Date: July 13, 1999

No Sequence report available

Sequence Info

GELS: GEL461

AA Analysis(mg/ml)

OD 280

Endotoxin Units/mg Protein

Prot A ppm

PUR Done Date

Yield Volume: 4.5ml

Cancel Reason

Status: Done

Origin

Storage Location

ASY | DNA | DOM | EXP | FAM | ELS | LIB | LQT | MAP | QLI | PRB | PRQ | PUR | RNA | SRC | UNQ | XPT | YST

Assay Viewer | Sequence Viewer | Gene Viewer | GenesGenes | SAGE

GenesGenes Feedback

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## **EXHIBIT D**

(3 pages; pages 29-31)

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EXHIBIT D—PAGE 1

GENENGINE'S

GENE VIEWER | SEQUENCE VIEWER | ASSAY VIEWER

GENE | TEAM | MAP | GENES | DNA | SPC | RNA | LB | FS | OUT | DON | EXP | PUR | LOT | ASY

LOT2552

Gene Info

Inventor/lot Name

Control

Initial Volume

Comments

Status

Date Entered

Date Cancelled

Cancel Reason

Registry

Available

Storage Location

Related Protein Lists

Lot

PNX

PRQ

PUR

LOT Name

Protein Form Name

LOT2552

DNA84663

PRO1825

PUR1009

PIN1308-1

Human CTRP3 Poly-His

Assay LOT Distribution

ASY	Status	Distribution Date	Result Date	Assay Name
ASY1	Retired	9/28/99	11/5/99	Heart Neonatal Hypertrophy
ASY8	Running	4/21/00	5/23/00	Endothelial cell proliferation
ASY9	Running	4/21/00	5/23/00	Inhibition of VEGF stimulated endothelial cell growth
ASY24	Retired	9/2/99	9/28/99	MLR - Stimulatory
ASY32	Retired	11/4/99	12/14/99	Guinea pig Proinflammatory activity [hairless]
ASY33	Running	12/10/01		Miscellaneous
ASY34	On Hold	9/14/99	11/8/99	Hu Venous Endothelial Cell c-fos Induction Assay
ASY51	Retired	11/4/99	12/14/99	Guinea pig Vascular Constrictor activity [hairless]

Additional Resources:

UNQ753 PRO1825 Human CTRP3 Poly-His DNA84663 PUR1009 EXP2247

PIN1308-1

some comment

August 5, 1999

Yes

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EXHIBIT D—PAGE 2

ASY64	Retired	11/4/99	12/14/99	Proinflammatory/PMN infiltrate
ASY67	Retired	9/2/99	9/28/99	MLR - Inhibitory
ASY68	On Hold	10/18/99	11/8/99	Hu Venous Endothelial Cell Ca Flux Assay
ASY74	Retired	9/28/99	11/8/99	Inhibition of Heart Neonatal Hypertrophy Induced by LIF+ET-1
ASY75	Retired	9/28/99	11/8/99	Enhancement of Heart Neonatal Hypertrophy Induced by LIF+
ASY100	Running	8/20/99		Endotoxin Level (LAL)
ASY103	Running	9/1/99		Protein Gel Analysis
ASY106	Retired	10/2/99	12/1/99	Glucose and FFA uptake in Differentiated Skeletal Muscle
ASY106	Retired	12/3/99	1/4/00	Glucose and FFA uptake in Differentiated Skeletal Muscle
ASY107	Running	11/16/99	1/4/00	Fetal hemoglobin induction in an erythroblastic cell line
ASY110	Retired	10/22/99	11/10/99	Chondrocytes Re-differentiation Assay
ASY110	Retired	12/1/99	4/5/00	Chondrocytes Re-differentiation Assay
ASY110	Retired	12/15/99	3/27/00	Chondrocytes Re-differentiation Assay
ASY110	Retired	5/2/00	8/18/00	Chondrocytes Re-differentiation Assay
ASY110	Retired	5/16/00	8/18/00	Chondrocytes Re-differentiation Assay
ASY111	Retired	10/22/99	11/10/99	Chondrocyte Proliferation Assay
ASY111	Retired	12/1/99	4/5/00	Chondrocyte Proliferation Assay
ASY111	Retired	12/15/99	3/27/00	Chondrocyte Proliferation Assay
ASY111	Retired	5/2/00	8/18/00	Chondrocyte Proliferation Assay
ASY111	Retired	5/16/00	8/18/00	Chondrocyte Proliferation Assay
ASY118	Retired	1/12/00	2/1/00	Inhibition of A -Peptide Binding to Factor VIIA
ASY119	Retired	1/12/00	2/1/00	Inhibition of A - Peptide Binding to Factor VIIIE
ASY128	Retired	5/5/00	6/20/00	Cytokine Release in Human Whole Blood
ASY129	Retired	5/16/00	8/18/00	Chondrocytes re-differentiation by Fluorescence
ASY130	Retired	5/16/00	8/18/00	Chondrocytes Proliferation by fluorescence
ASY132	Retired	6/23/00	8/7/00	Activation of NFkB
ASY134	Retired	10/13/00	11/30/00	Activation of NIKB [ Luciferase]
ASY134	Retired	12/5/00	1/22/01	Activation of NIKB [ Luciferase]
ASY135	Retired	9/12/00	10/19/00	Induction of E-selectin
ASY138	Running	2/23/01	4/9/01	Normal Human Iliac Artery Endothelial cells

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ASY139	Running	2/23/01	4/9/01	Pooled Human Umbilical vein Endothelial cells
ASY140	Running	2/23/01	4/9/01	Coronary artery Smooth Muscle cells
ASY141	Running	2/23/01	4/9/01	Normal human Dermal Fibroblast Proliferation
ASY142	Running	2/14/01	3/26/01	NF-kappa B Inhibition Assay
ASY142	Running	3/8/01	3/26/01	NF-kappa B Inhibition Assay
ASY146	Running	7/19/01	8/3/01	Human Microvascular Endothelial Cell Proliferation Assay
ASY162	Running	11/16/99	9/5/00	NCI Oncology Screen-1
ASY165	Running	8/1/01	9/19/01	CREB
ASY165	Running	9/19/01	9/24/01	CREB
ASY170	Piloting	11/9/01	11/16/01	NHEK proliferation assay
ASY174	Piloting	3/12/02	4/3/02	Bovine Retinal M Endothelial
ASY174	Piloting	4/4/02		Bovine Retinal M Endothelial
ASY174	Piloting	5/17/02		Bovine Retinal M Endothelial
ASY174	Piloting	11/20/02		Bovine Retinal M Endothelial
ASY175	Running	12/21/01		Neuronal Differentiation using Rinat technology
ASY175	Running	5/30/02		Neuronal Differentiation using Rinat technology
ASY176	Piloting	5/31/02		Hemoglobin Assay
ASY176	Piloting	7/16/02		Hemoglobin Assay
ASY177	Piloting	4/22/03	8/18/03	fibroblast migration assay
ASY178	Running	1/23/03		Proliferation of Fibroblasts
ASY180	Running	3/11/03	3/25/03	Mouse Keratinocyte Assay
ASY181	Running	3/6/03	3/13/03	Human Mammary Epithelial Cell Assay

ASY | DNA | DOM | EXP | FAM | ELS | LIB | LOT | MAP | LOU | PER | PRQ | PUR | RNA | SEC | UNQ | XPT | YST  
 Assay Viewer | Sequence Viewer | Gene Viewer | GeneGator | SAGE

GeneGator Feedback



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## **EXHIBIT E**

(2 pages; pages 33-34)

EXHIBIT E—PAGE 1

### GreenGated Feedback

EXHIBIT E—PAGE 2

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